## AMENDMENT TO THE CLAIMS:

The current listing of claims as follows will replace all previously entered claims. :

## **Listing of Claims**

- 1. (Currently Amended) An Fe variant antibody and/or immunoadhesin of a parent Fc polypeptide, said Fe variant antibody and/or immunoadhesin comprising at least one amino acid modification in the Fc region of said parent Fc polypeptide said modification occurring at a position selected from the group consisting of: 228, 230, 231, 232, 240, 244, 245, 247, 262, 263, 266, 271, 273, 275, 281, 284, 291, 299, 302, 304, 313, 323, 325, 328, 332, and 336, wherein said Fe variant antibody and/or immunoadhesin protein exhibits altered binding to an FcγR as compared to the parent Fc polypeptide, wherein numbering is according to the EU index.
- 2. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 1, wherein said Fe variant antibody and/or immunoadhesin comprises at least one substitution selected from the group consisting of: 230, 240, 244, 245, 247, 262, 263, 266, 273, 275, 299, 302, 313, 323, 325, 328, and 332.
- 3. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 1, wherein said altered binding is an increase in affinity of said Fe variant antibody and/or immunoadhesin to said FeyR.
- 4. (Currently Amended) An Fe-variant antibody and/or immunoadhesin according to claim 3, wherein said Fe variant antibody and/or immunoadhesin binds with greater affinity to a mouse FcγR.
- 5. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 3, wherein said said FcγR is a human Fc receptor selected from the group consisting of FcγRI, FcγRIIa, FcγRIIb, FcγRIIc, and FcγRIIIa.

- 6. (Currently Amended) An Fe-variant antibody and/or immunoadhesin according to claim 3, wherein said Fe-variant antibody and/or immunoadhesin binds with greater affinity to human FcγRI and FcγRIIIa, but exhibits unaltered affinity to a human receptor selected from the group consisting of FcγRIIa, FcγRIIb, and FcγRIIc.
- 7. (Currently Amended) Fe variant antibody and/or immunoadhesin according to claim 6, wherein said Fe variant antibody and/or immunoadhesin exhibits unaltered affinity to FcyRIIa, FcyRIIb, and FcyRIIc.
- 8. (Withdrawn currently amended) An Fe variant antibody and/or immunoadhesin according to claim 3, wherein the affinity increase for binding to one or more human FcγRII's is greater than the affinity increase for binding to human FcγRII or FγRIIIa.
- 9. (Withdrawn currently amended) An Fe variant antibody and/or immunoadhesin according to claim 3, wherein the affinity increase is the same for binding to FcγRIIa, FcγRIIb, and FcγRIIc.
- 10. (Withdrawn currently amended) An Fe-variant antibody and/or immunoadhesin according to claim 3, wherein the affinity increase for binding to FcγRIIc is greater than the affinity increase for binding to FcγRIIb.
- 11. (Withdrawn) An Fe variant antibody and/or immunoadhesin according to claim 3, wherein said Fe variant antibody and/or immunoadhesin binds with greater affinity to human FcγRIIa, but exhibits unaltered affinity to a human receptor selected from the group consisting of FcγRI, FcγRIIb, FcγRIIc, and FcγRIIIa.
- 12. (Withdrawn currently amended) An Fe-variant antibody and/or immunoadhesin according to claim 11, wherein said Fe-variant antibody and/or immunoadhesin exhibits unaltered affinity to FcγRI, FcγRIIb, FcγRIIc, and FcγRIIIa.

13. (Currently Amended) An Fe-variant antibody and/or immunoadhesin according to any of claims 1–12 1-7, wherein said Fe variant antibody and/or immunoadhesin binds with greater affinity to FeRn.

- 14. (Withdrawn currently amended) An Fe variant antibody and/or immunoadhesin according to any of claims 1-12, wherein said Fe variant antibody and/or immunoadhesin binds with unaltered affinity to FcRn.
- 15. (Withdrawn currently amended) An Fe-variant antibody and/or immunoadhesin according to any of claims 1-12, wherein said Fe variant antibody and/or immunoadhesin binds with reduced affinity to FcRn.
- 16. (Withdrawn currently amended) An Fe variant antibody and/or immunoadhesin according to any of claims 1-12, wherein said Fe variant antibody and/or immunoadhesin binds with greater affinity to C1q or enhances complement dependent cytotoxicity (CDC).
- 17. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to any of claims 1–12 1-7, wherein said Fe variant antibody and/or immunoadhesin binds with unaltered affinity to C1q or enhances CDC.
- 18. (Withdrawn currently amended) An Fe variant antibody and/or immunoadhesin according to claim 1, wherein said Fe variant antibody and/or immunoadhesin further comprises an engineered glycoform.
- 19. (Withdrawn currently amended) An Fe variant antibody and/or immunoadhesin according to claim 18 wherein said engineered glycoform comprises an altered level of fucosylation or bisecting oligosaccharides as compared to the parent Fc polypeptide.
- 20. (Withdrawn) An Fe variant antibody and/or immunoadhesin according to claim 18 wherein said engineered glycoform improves effector function.

21. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 1, wherein said Fe variant antibody and/or immunoadhesin enhances or reduces an effector function selected from the group consisting of antibody dependent cell-mediated cytotoxicity (ADCC), CDC, or antibody dependent cell-mediated phagocytosis (ADCP)\_function as compared to the parent Fc polypeptide.

- 22. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 21, wherein said effector function is antibody dependent cell-mediated cytotoxicity (ADCC).
- 23. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 22, wherein said Fe variant antibody and/or immunoadhesin improves ADCC in the presence of human effector cells as compared to said parent Fc polypeptide.
- 24. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 22, wherein said ADCC improvement is an enhancement in potency such that the EC50 of said Fe variant antibody and/or immunoadhesin is approximately 5-fold greater than that of said parent Fc polypeptide.
- 25. (Currently Amended) An Fe-variant antibody and/or immunoadhesin according to claim 23, wherein said ADCC improvement is an enhancement in potency such that the EC50 of said Fe-variant antibody and/or immunoadhesin is between approximately 5-fold and 1000-fold greater than that of said parent Fc polypeptide.
- 26. (Currently Amended) An Fe-variant antibody and/or immunoadhesin according to claim 23, wherein said ADCC improvement is an enhancement in efficacy such that the maximal ADCC is approximately 2-fold greater than that of said parent Fc polypeptide.

- 27. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 26, wherein said Fe variant antibody and/or immunoadhesin improves ADCC in the presence of mouse effector cells as compared to said parent Fe polypeptide.
- 28. (Currently Amended) An Fe-variant antibody and/or immunoadhesin according to claim 21, wherein said effector function is ADCP.
- 29. (Currently Amended) An Fe-variant antibody and/or immunoadhesin according to claim 28, wherein said Fe-variant antibody and/or immunoadhesin improves ADCP as compared to said parent Fc polypeptide.
- 30. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 25 or 29, wherein CDC is unaffected.
- 31. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 25 or 29, wherein CDC is ablated.
- 32. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 1 wherein said polypeptide has specificity for a target antigen selected from the group consisting of CD4, CD19, CD20, CD22, CD25, CD30, CD33, CD52, CD80, B7-1, B7-2, CLTA-4, BAFF-R, Her2/neu, EGFR, EpCAM, MUC1, GD3, CEA, CA 125, HLA-DR, TNFalpha, and VEGF.
- 33. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 1 or 21 comprising a modification selected from the group consisting of 228E, 228K, 228Y, 228G, 230E, 230Y, 230G, 231E, 231K, 231Y, 231P, 231G, 232E, 232K, 232Y, 232G, 262E, 262F, 271D, 271E, 271N, 271Q, 271K, 271R, 271S, 271T, 271H, 271A, 271V, 271L, 271I, 271F, 271M, 271Y, 271W, 271G, 275L, 281D, 281K, 281Y, 281P, 284E, 284N, 284T, 284L, 284Y, 291D, 291E, 291Q, 291T, 291H, 291I, 291G, 299D, 299E, 299N, 299Q, 299K, 299R,

299L, 299F, 299M, 299Y, 299W, 299P, 299G, 304D, 304N, 304T, 304H, 304L, 325K, 325R, 325S, 325F, 325M, 325Y, 325W, 325P, 325G, 328D, 328Q, 328K, 328R, 328S, 328T, 328V, 328I, 328Y, 328W, 328P, 328G, 332K, 332R, 332S, 332V, 332L, 332F, 332M, 332W, 332P, 332G, 336E, 336K, 336Y, 230A, 240A, 240I, 240M, 240T, 244H, 245A, 247G, 247V, 262A, 262E, 262I, 262T, 263A, 263I, 263M, 263T, 266A, 266I, 266M, 266T, 273I, 275W, 299A, 299E, 299F, 299H, 299I, 299S, 299V, 302I, 313F, 323I, 325A, 325D, 325E, 325H, 325I, 325L, 325Q, 325T, 325V, 328A, 328D, 328E, 328F, 328H, 328I, 328M, 328N, 328Q, 328T, 328V, 332A, 332D, 332E, 332H, 332N, 332Q, 332T, and 332Y.

- 34. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 1 wherein said parent Fc polypeptide is an antibody comprising said Fe variant antibody and/or immunoadhesin.
- 35. (Currently Amended) An Fe variant antibody and/or immunoadhesin according to claim 1 wherein said Fe variant antibody and/or immunoadhesin is an Fc fusion protein comprising said Fe variant antibody and/or immunoadhesin.
- 36. (Currently Amended) A pharmaceutical composition comprising an Fe variant antibody and/or immunoadhesin according to claim 1 and a pharmaceutically acceptable carrier.
- 37. (Withdrawn currently amended) A method of treating a mammal in need of said treatment, comprising administering a variant antibody and/or immunoadhesin protein of claim 1.

38. (Withdrawn – currently amended) An Fe variant antibody and/or immunoadhesin according to claims 1 or 33 comprising at least one modification selected from the group consisting of A330L/I332E, D265F/N297E/I332E, D265Y/N297D/I332E,

D265Y/N297D/T299L/I332E, F241E/F243Q/V262T/V264E,

F241E/F243Q/V262T/V264E/I332E, F241E/F243R/V262E/V264R,

F241E/F243R/V262E/V264R/I332E, F241E/F243Y/V262T/V264R,

F241E/F243Y/V262T/V264R/I332E, F241L/F243L/V262I/V264I, F241L/V262I,

F241R/F243Q/V262T/V264R, F241R/F243Q/V262T/V264R/I332E,

F241W/F243W/V262A/V264A, F241Y/F243Y/V262T/V264T,

F241Y/F243Y/V262T/V264T/N297D/I332E, F243L/V262I/V264W, F243L/V264I,

L328D/I332E, L328E/I332E, L328H/I332E, L328I/I332E, L328M/I332E, L328N/I332E,

L328Q/I332E, L328T/I332E, L328V/I332E, N297D/A330Y/I332E, N297D/I332E,

N297D/I332E/S239D/A330L, N297D/S298A/A330Y/I332E, N297D/T299E/I332E,

N297D/T299F/I332E, N297D/T299H/I332E, N297D/T299I/I332E, N297D/T299L/I332E,

N297D/T299V/I332E, N297E/I332E, N297S/I332E, P230A/E233D/I332E,

P244H/P245A/P247V, S239D/A330L/I332E, S239D/A330Y/I332E,

S239D/A330Y/I332E/K326E, S239D/A330Y/I332E/K326T, S239D/A330Y/I332E/L234I,

S239D/A330Y/I332E/L235D, S239D/A330Y/I332E/V240I, S239D/A330Y/I332E/V264T,

S239D/A330Y/I332E/V266I, S239D/D265F/N297D/I332E, S239D/D265H/N297D/I332E,

S239D/D265I/N297D/I332E, S239D/D265L/N297D/I332E, S239D/D265T/N297D/I332E,

\$239D/D265V/N297D/I332E, \$239D/D265Y/N297D/I332E, \$239D/I332D, \$239D/I332E,

S239D/I332E/A330I, S239D/I332N, S239D/I332Q, S239D/N297D/I332E,

S239D/N297D/I332E/A330Y, S239D/N297D/I332E/A330Y/F241S/F243H/V262T/V264T,

S239D/N297D/I332E/K326E, S239D/N297D/I332E/L235D, S239D/S298A/I332E,

S239D/V264I/A330L/I332E, S239D/V264I/I332E, S239D/V264I/S298A/I332E, S239E/D265N,

S239E/D265O, S239E/I332D, S239E/I332E, S239E/I332N, S239E/I332O,

S239E/N297D/I332E, S239E/V264I/A330Y/I332E, S239E/V264I/I332E,

\$239E/V264I/\$298A/A330Y/I332E, \$239N/A330L/I332E, \$239N/A330Y/I332E,

S239N/I332D, S239N/I332E, S239N/I332N, S239N/I332Q, S239N/S298A/I332E,

S239Q/I332D, S239Q/I332E, S239Q/I332N, S239Q/I332Q, S239Q/V264I/I332E, S298A/I332E,

V264E/N297D/I332E, V264I/A330L/I332E, V264I/A330Y/I332E, V264I/I332E, V264I/S298A/I332E, Y296D/N297D/I332E, Y296E/N297D/I332E, Y296H/N297D/I332E, Y296N/N297D/I332E, Y296Q/N297D/I332E, and Y296T/N297D/I332E.

39. (Withdrawn - currently amended) An Fe variant antibody and/or immunoadhesin according to claims 1, 33 or 38 further comprising a modification selected from the group consisting of 227E, 227K, 227Y, 227G, 233N, 233Q, 233K, 233R, 233S, 233T, 233H, 233A, 233V, 233L, 233I, 233F, 233M, 233Y, 233W, 233G, 234K, 234R, 234S, 234A, 234M, 234W, 234P, 234G, 235E, 235K, 235R, 235A, 235M, 235W, 235P, 235G, 236D, 236E, 236N, 236Q, 236K, 236R, 236S, 236T, 236H, 236A, 236V, 236L, 236I, 236F, 236M, 236Y, 236W, 236P, 237D, 237E, 237N, 237Q, 237K, 237R, 237S, 237T, 237H, 237V, 237L, 237I, 237F, 237M, 237Y, 237W, 237P, 238D, 238E, 238N, 238Q, 238K, 238R, 238S, 238T, 238H, 238V, 238L, 238I, 238F, 238M, 238Y, 238W, 238G, 239Q, 239K, 239R, 239V, 239L, 239I, 239M, 239W, 239P, 239G, 241D, 241E, 241Y, 243E, 246D, 246E, 246H, 246Y, 249Q, 249H, 249Y, 255E, 255Y, 258S, 258H, 258Y, 260D, 260E, 260H, 260Y, 264D, 264E, 264N, 264Q, 264K, 264R, 264S, 264H, 264W, 264P, 264G, 265Q, 265K, 265R, 265S, 265T, 265H, 265V, 265L, 265I, 265F, 265M, 265Y, 265W, 265P, 267E, 267Q, 267K, 267R, 267V, 267L, 267I, 267F, 267M, 267Y, 267W, 267P, 268D, 268E, 268Q, 268K, 268R, 268T, 268V, 268L, 268I, 268F, 268M, 268W, 268P, 268G, 269K, 269S, 269V, 269I, 269M, 269W, 269P, 269G, 270R, 270S, 270L. 270I, 270F, 270M, 270Y, 270W, 270P, 270G, 272D, 272R, 272T, 272H, 272V, 272L, 272F, 272M, 272W, 272P, 272G, 274D, 274N, 274S, 274H, 274V, 274I, 274F, 274M, 274W, 274P, 274G, 276D, 276T, 276H, 276V, 276I, 276F, 276M, 276W, 276P, 276G, 278D, 278N, 278Q, 278R, 278S, 278H, 278V, 278L, 278I, 278M, 278P, 278G, 280K, 280L, 280W, 280P, 280G, 282E, 282K, 282Y, 282P, 282G, 283K, 283H, 283L, 283Y, 283P, 283G, 285D, 285E, 285O, 285K, 285Y, 285W, 286E, 286Y, 286P, 286G, 288D, 288E, 288Y, 290D, 290N, 290H, 290L, 290W, 292D, 292E, 292T, 292Y, 293N, 293R, 293S, 293T, 293H, 293V, 293L, 293I, 293F, 293M, 293Y, 293W, 293P, 293G, 294K, 294R, 294S, 294T, 294H, 294V, 294L, 294I, 294F, 294M, 294Y, 294W, 294P, 294G, 295D, 295E, 295N, 295R, 295S, 295T, 295H, 295V, 295L 295F, 295M, 295Y, 295W, 295P, 295G, 296K, 296R, 296A, 296V, 296M, 296G, 297O, 297K, 297R, 297T, 297H, 297V, 297L, 297I, 297F, 297M, 297Y, 297W, 297P, 297G, 298D, 298E,

298O, 298K, 298R, 298I, 298F, 298M, 298Y, 298W, 300D, 300E, 300N, 300Q, 300K, 300R, 300S, 300T, 300H, 300A, 300V, 300M, 300W, 300P, 300G, 301D, 301E, 301H, 301Y, 303D, 303E, 303Y, 305E, 305T, 305Y, 317E, 317Q, 318Q, 318H, 318L, 318Y, 320N, 320S, 320H, 320V, 320L, 320F, 320Y, 320W, 320P, 320G, 322D, 322S, 322V, 322I, 322F, 322Y, 322W, 322P, 322G, 324H, 324F, 324M, 324W, 324P, 324G, 326P, 327E, 327K, 327R, 327H, 327V, 327I, 327F, 327M, 327Y, 327W, 327P, 329D, 329E, 329N, 329Q, 329K, 329R, 329S, 329T, 329H, 329V, 329L, 329I, 329M, 329Y, 329W, 329G, 330E, 330N, 330T, 330P, 330G, 331D, 331O, 331R, 331T, 331L, 331I, 331F, 331M, 331Y, 331W, 333L, 333F, 333M, 333P, 334P, 335N, 335S, 335H, 335V, 335L, 335I, 335F, 335M, 335W, 335P, 335G, 337E, 337N, 337H, 233D, 234D, 234E, 234F, 234H, 234I, 234N, 234Q, 234T, 234V, 234Y, 235D, 235F, 235H, 235I, 235N, 235Q, 235S, 235T, 235V, 235Y, 239D, 239E, 239F, 239H, 239N, 239Q, 239T, 239Y, 241E, 241L, 241R, 241S, 241W, 241Y, 243H, 243L, 243Q, 243R, 243W, 243Y, 264A, 264E. 264F. 264I. 264L. 264M. 264R. 264T. 264W. 264Y. 265F. 265G. 265H. 265I. 265L. 265N, 265O, 265T, 265V, 265Y, 267D, 267H, 267L, 267N, 267Q, 267T, 269F, 269H, 269L, 269N, 269R, 269T, 269Y, 270H, 270O, 270T, 272I, 272K, 272S, 272Y, 274E, 274L, 274R, 274T, 274Y, 276E, 276L, 276R, 276S, 276Y, 278E, 278K, 278T, 278W, 283R, 296D, 296E, 296H, 296L, 296L, 296N, 296O, 296S, 296T, 297D, 297E, 297S, 298A, 298H, 298N, 298T, 318R, 320D, 320I, 320T, 322H, 322T, 324D, 324I, 324L, 324R, 324T, 324V, 324Y, 326E, 326I, 326L, 326T, 327D, 327L, 327N, 327S, 327T, 329F, 330F, 330H, 330L, 330L, 330M, 330R, 330S, 330V, 330W, 330Y, 331H, 331V, 333A, 333H, 333I, 333T, 333Y, 334A, 334F, 334I, 334T, 335D, 335R, and 335Y.

- 40. (Withdrawn currently amended) An Fe variant antibody and/or immunoadhesin according to claim 1 wherein said Fe variant antibody and/or immunoadhesin further comprises a set of modifications selected from 241W/243W, 267Q/327S, 243L/264I, 234I/235D, 264E/297D/332E, 239E/265N, 239E/265Q, 239E/265G, 267L/327S and 240I/266I.
- 41. (Withdrawn currently amended) An Fe variant antibody and/or immunoadhesin according to claim 33 wherein said antibody is a full length antibody.

42. (Withdrawn – currently amended) An Fe variant antibody and/or immunoadhesin according to claim 33 wherein said antibody is a human antibody.

43. (Withdrawn – currently amended) An Fe variant antibody and/or immunoadhesin according to claim 33 wherein said antibody is an antibody fragment.

44-45. (Canceled)